PCT

WORLD INTELLECTUAL PROPERTY ORGANIZATION International Bureau



INTERNATIONAL APPLICATION PUBLISHED UNDER THE PATENT COOPERATION TREATY (PCT)

(51) International Patent Classification 6:

(11) International Publication Number:

WO 97/36996

C12N 1/12, 1/14, C12P 7/64, C12N 1/00

(43) International Publication Date:

9 October 1997 (09.10.97)

(21) International Application Number:

PCT/EP97/01446

A3

(22) International Filing Date:

21 March 1997 (21.03.97)

(30) Priority Data:

96200837.1

28 March 1996 (28.03.96)

EP

(34) Countries for which the regional or international application was filed:

NL et al.

(71) Applicant (for all designated States except US): GIST-BROCADES B.V. [NL/NL]; Wateringseweg I, P.O. Box 1, NL-2600 MA Delft (NL).

(72) Inventors; and

(75) Inventors/Applicants (for US only): BIJL, Hendrik, Louis [NL/NL]; Insulindestraat 72, NL-3131 ZD Vlaardingen (NL). SCHAAP, Albert [NL/NL]; Smitshoek 13, NL-2993 BG Barendrecht (NL). VISSER, Johannes, Martinus, Jacobus [NL/NL]; Gangboord 102, NL-3823 TJ Amersfoort (NL).

(74) Agents: GOLDIN, Douglas, Michael et al.; J.A. Kemp & Co., 14 South Square, Gray's Inn, London WC1R 5LX (GB).

(81) Designated States: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GE, GH, HU, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ARIPO patent (GH, KE, LS, MW, SD, SZ, UG), Eurasian patent (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European patent (AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE), OAPI patent (BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG).

Published

With international search report.

Before the expiration of the time limit for amending the claims and to be republished in the event of the receipt of amendments.

(88) Date of publication of the international search report: 22 January 1998 (22.01.98)

(54) Title: PROCESS FOR THE PREPARATION OF A GRANULAR MICROBIAL BIOMASS AND ISOLATION OF VALUABLE COMPOUNDS THEREFROM

(57) Abstract

A process for the isolation of desired compound(s) from a microbial biomass is diclosed, wherein the microbial biomass (which, if necessary, is pretreated to give a dry matter content of from 25 to 80 %) is granulated (e.g. by extrusion) and then dried to a dry matter content of at least 80 %. The granulation of the biomass to granules significantly eases subsequent drying of the biomass (which can be stored as dried granules) and gives higher yields on extraction of the compound(s).

FOR THE PURPOSES OF INFORMATION ONLY

Codes used to identify States party to the PCT on the front pages of pamphlets publishing international applications under the PCT.

AL	Albania	ES	Spain	I.S	Lesotho		
M	Armenia	FI	Finland	LT	Lithuania	SI	Slovenia
LT.	Austria	FR	Prance	LU		SK	Slovakia
U	Australia	GA	Gabon	LV	Luxembourg	SN	Senegal
Z	Azerbaijan	GB	United Kingdom	MC	Latvia	SZ	Swaziland
A	Bosnia and Herzegovina	GE	Georgia	MD	Monaco	TD	Chad
B	Barbados	GH	Ghana		Republic of Moldova	TG	Togo
E	Belgium	GN	Guinea	MG	Madagascar	TJ	Tajikistan
F	Burkina Faso	GR	Greece	MK	The former Yugoslav	TM	Turkmenistan
G	Bulgaria	HU	Hungary	141	Republic of Macedonia	TR	Turkey
IJ	Benin	IE	ireland	ML	Mali	TT	Trinidad and Tobago
BR	Brazil	iL	Israel	MN	Mongolia	UA	Ukraine
BY	Belarus	is	Iceland	MR	Mauritania	UG	Uganda
CA	Canada	IT	Italy	MW	Malawi	US	United States of America
CF	Central African Republic	JP	Japan	MX NE	Mexico	UZ	Uzbekistan
CG	Congo	KE	Kenya	NL NL	Niger	VN	Viet Nam
CH	Switzerland	KG	Kyrgyzstan	NO NO	Netherlands	YU	Yugoslavia
CI	Côte d'Ivoire	KP	Democratic People's	NZ NZ	Norway	zw	Zimbabwe
CM	Cameroon		Republic of Korea	PL	New Zealand		
CN	China	KR	Republic of Korea	PT	Poland		
CU	Cuba	KZ	Kazakatan	RO	Portugal		
CZ.	Czech Republic	LC	Saint Lucia	RU	Romania		
E	Germany	u	Licchtenstein	SD	Russian Federation		
K	Denmark	LK	Sri Lanka	SE	Sudan Sweden		
Œ	Estonia	LR	Liberia	SG.	Singapore		

Intern 181 Application No PCT/EP 97/01446

A. CLASSIFICATION OF SUBJECT MATTER
IPC 6 C12N1/12 C12N1/14 C12N1/00 C12P7/64 According to International Patent Classification (IPC) or to both national classification and IPC 8. FIELDS SEARCHED Minimum documentation searched (classification system followed by classification symbols) C12N C12P IPC 6 Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched Electronic data base consulted during the international search (name of data base and, where practical, search terms used) C. DOCUMENTS CONSIDERED TO BE RELEVANT Relevant to olsim No. Citation of document, with indication, where appropriate, of the relevant passages Category ° 1-7,30, GB 1 466 853 A (SIMON ROSEDOWNS LTD) 1974 X 34-36 cited in the application 8-24 see the whole document Y 1-7,30, FR 2 210 662 A (BRITISH PETROLEUM CO) 12 Х 34-36 July 1974 see the whole document 1-7,30, DD 150 627 A (AKADEMIE DER WISSENSCHAFTEN X 34-36 DER DDR) 1981 see the whole document 1-7. DE 19 23 529 A (VEBINGENIEURTECHNISCHES X 34-36 ZENTRALBURO MINERALÖLE U. ORGANISCHE GRUNSDTOF) 1969 see the whole document -/--Patent family members are listed in annex. X Further documents are listed in the continuation of box C. Special categories of cited documents : "I" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the "A" document defining the general state of the art which is not considered to be of particular relevance "E" earlier document but published on or after the international "X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to filing date involve an inventive step when the document is taken alone "L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified) "Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such docu-ments, such combination being obvious to a person skilled *O* document referring to an oral disclosure, use, exhibition or "P" document published prior to the international filing date but later than the priority date claimed "&" document member of the same patent family Date of mailing of the international search report Date of the actual completion of the international search 0 2 -12 - 1997 28 October 1997 Authorized office: Name and mailing address of the ISA European Patent Office, P.B. 5818 Patentiaan 2 NL - 2280 HV Riswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl, Gurdjian, D Fax: (+31-70) 340-3016

4

Intern. :al Application No PCT/EP 97/01446

	ation) DOCUMENTS CONSIDERED TO BE RELEVANT	PCT/EP 97/01446
Category °	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to plaim No.
Υ	JAREONKITMONGKOL S ET AL: "PRODUCTION OF DIHOMO-GAMMA-LINOLENIC ACID BY A DELTA-5 DESATURASE-DEFECTIVE MUTANT OF MORTIERELLA-ALPINA 1S-4" APPLIED AND ENVIRONMENTAL MICROBIOLOGY, 58 (7). 1992. 2196-2200., XP000676362 see the whole document	8-11, 15-18
Y	JAREONKITMONGKOL S ET AL: "PRODUCTION OF AN EICOSAPENTAENOIC ACID-CONTAINING OIL BY A DELTA-12 DESATURASE-DEFECTIVE MUTANT OF MORTIERELLA-ALPINA 1S-4" JOURNAL OF THE AMERICAN OIL CHEMISTS' SOCIETY, 70 (2). 1993. 119-123., XP000676339 see the whole document	8-11, 15-18
	WO 94 29292 A (KRKA TOVARNA ZDRAVIL ;HAJKO PAVICA (SI); VESEL TANJA (SI); RADEZ I) 22 December 1994 see claims 1-8	19,20
	WO 93 20183 A (ZEAGEN INC) 14 October 1993 see claims 1-35,71-97	19,20
	DATABASE WPI Week 9210 Derwent Publications Ltd., London, GB; AN 92-077323 XP002035614 & JP 04 023 979 A (KOBE STEEL LTD), 28 January 1992 see abstract	23,24
	EP 0 688 871 A (QUEST INT) 27 December 1995 see abstract	21,22
	WO 92 12711 A (MARTEK CORP) 6 August 1992 see claims 1-25	12-14
	WO 91 11918 A (MARTEK CORP) 22 August 1991 see abstract; claims 1-55	34,35
	EP 0 322 227 A (SUNTORY LTD) 28 June 1989	9-11, 15-18
	see the whole document	
	US 4 056 638 A (HUANG HUA-FENG ET AL) 1 November 1977 see abstract; figures 2-5 see column 5, line 58 - column 6, line 5; claim 1; examples 1-4	27,29-32 28
- 1	-/	1

4

Interr all Application No PCT/EP 97/01446

0.10	ALLAN DOCUMENTO CONGIDERES TO BE RELEVANT	
	ation) DOCUMENTS CONSIDERED TO BE RELEVANT Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
Category *	OMBINITY OF GOODINALITY AND INCIDENTAL AND A STATE OF THE ISSUED PROPERTY.	
Y	KYLE D J; RATLEDGE C ED.: "Industrial applications of single cell oils." 1992 , AMERICAN OIL CHEMISTS SOCIETY , CHAMPAIGN XP002044902	28
Α	see page 61; figures 5-14	10,15
A	see page 61; figures 5-14	10,15
	·	

Form PCT/ISA/210 (continuation of second sheet) (July 1992)

rnational application No.

PCT/EP 97/01446

Boxi	Observations where certain claims were found unsearchable (Continuation of item 1 of first sheet)
This Inte	ernational Search Report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:
1.	Claims Nos.: because they relate to subject matter not required to be searched by this Authority, namely:
	Claims Nos.: because they relate to parts of the International Application that do not comply with the prescribed requirements to such an extent that no meaningful International Search can be carried out, specifically:
	Claims Nos.: because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a)
Box (I	Observations where unity of invention is lacking (Continuation of Item 2 of first sheet)
This Intern	national Searching Authority found multiple inventions in this international application, as follows:
	See annex
1. X A	is all required additional search fees were timely paid by the applicant, this International Search Report covers all earchable claims.
2. A	s all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment I any additional fee.
3. A	s only some of the required additional search fees were timely paid by the applicant, this International Search Report overs only those claims for which fees were paid, specifically claims Nos
4. No re	o required additional search fees were timely paid by the applicant. Consequently, this International Search Report is stricted to the invention first mentioned in the claims; it is covered by claims Nos.:
Remark on	Protest X The additional search fees were accompanied by the applicant's protest. No protest accompanied the payment of additional search fees.

FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

Invention 1: claims 1-26, 34-36

processes for the isolation of compounds from a microbial biomass ,the process comprising obtaining a biomass with a dry matter content from 25%-80%, granulating these particles with a dry matter content from 25%-80%, drying the the granules to a dry dry matter content of at least 80%, purification, extraction or isolation of the compound and use of the extracted compounds in food or cosmetic/pharmaceutical compositions.

Invention 2: claims 27-33

Microbial extrudates or microbial biomass granular compositions having an average of 30-70% dry matter- or at least 80% dry matter

information on patent family members

Intern: at Application No PCT/EP 97/01446

			1/EP 9//01446
Patent document cited in search report	Publication date	Patent family member(s)	Publication date
GB 1466853 A	09-03-77	DE 2424612 A FR 2230388 A NL 7406900 A,	30-04-75 20-12-74 B, 26-11-74
FR 2210662 A	12-07-74	GB 1411450 A	22-10-75
DD 150627 A		NONE	
DE 1923529 A	04-12-69	NONE	
WO 9429292 A	22-12-94	SI 9300303 A BG 100204 A CA 2164411 A CZ 9503251 A EP 0702679 A HU 72836 A PL 311880 A SK 148095 A	31-12-94 28-06-96 22-12-94 17-04-96 27-03-96 28-05-96 18-03-96 03-07-96
WO 9320183 A	14-10-93	AU 3919193 A US 5422247 A	08-11-93 06-06-95
EP 0688871 A	27-12-95	JP 8168392 A US 5618706 A	02-07-96 08-04-97
WO 9212711 A	06-08-92	AU 661297 B AU 1239292 A BR 9205526 A CA 2101274 A EP 0568606 A IL 100733 A IL 114253 A JP 6505153 T MX 9200320 A NZ 241359 A US 5550156 A US 5374657 A	20-07-95 27-08-92 19-04-94 25-07-92 10-11-93 31-12-95 13-07-97 16-06-94 01-10-92 26-08-94 27-08-96 20-12-94
√O 9111918 A	22-08-91	US 5407957 A AU 660162 B	18-04-95 15-06-95

information on patent family members

Intern ini Application No
PCT/EP 97/01446

Patent document cited in search report	Publication date	Patent family member(s)	Publication date
WO 9111918 A		AU 7330291 A CA 2076018 A EP 0515460 A IL 97126 A JP 5503425 T US 5492938 A US 5397591 A	03-09-91 14-08-91 02-12-92 15-03-95 10-06-93 20-02-96 14-03-95
EP 0322227 A	28-06-89	JP 1243992 A AT 124458 T CA 1316861 A DE 3854080 D DE 3854080 T ES 2074053 T US 4916066 A	28-09-89 15-07-95 27-04-93 03-08-95 02-11-95 01-09-95 10-04-90
US 4056638 A	01-11-77	BR 7603832 A CA 1070159 A DE 2626265 A FR 2314940 A GB 1496113 A JP 1248024 C JP 52001082 A JP 59023771 B SE 7605616 A	05-04-77 22-01-80 30-12-76 14-01-77 30-12-77 16-01-85 06-01-77 05-06-84 17-12-76